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**PROVISIONAL APPLICATION FOR PATENT COVER SHEET**  
This is a request for filing a PROVISIONAL APPLICATION FOR PATENT under 37 CFR 1.53(c).

**INVENTOR(S)**

Given Name (first and middle [if any])	Family Name or Surname	Residence (City and either State or Foreign Country)
Dieter	Manstein	Boston, Massachusetts

Additional inventors are being named on the 1 separately numbered sheets attached hereto

**TITLE OF THE INVENTION (280 characters max)**

METHOD AND APPARATUS FOR DERMATOLOGICAL TREATMENT AND FRACTIONAL SKIN RESURFACING

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**ENCLOSED APPLICATION PARTS (check all that apply)**

Specification Number of Pages

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Other (specify)

Claims - No. of sheets 8  
Abstract - No. of sheets 1

Application Data Sheet. See 37 CFR 1.76

**METHOD OF PAYMENT OF FILING FEES FOR THIS PROVISIONAL APPLICATION FOR PATENT**

Applicant claims small entity status. See 37 CFR 1.27.

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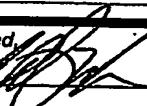
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The invention was made by an agency of the United States Government or under a contract with an agency of the  
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Respectfully submitted,

SIGNATURE 

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Date: March 27, 2003

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# BAKER BOTT

**PROVISIONAL APPLICATION COVER SHEET**  
*Additional Page*

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BAKER BOTTS, L.L.P.

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TO WHOM IT MAY CONCERN:

Be it known that we, Dieter Manstein and Richard Anderson, citizens of Germany and the United States of America, whose post office addresses are 1 Longfellow Place #2624, Boston, MA 02114 and 399 Marrett Road, Lexington, MA 02421, respectively, have made an invention in a

**METHOD AND APPARATUS FOR DERMATOLOGICAL TREATMENT AND  
FRACTIONAL SKIN RESURFACING**

of which the following is a

**SPECIFICATION**

**BACKGROUND OF THE INVENTION**

**1. Field Of The Invention**

[0001] The present invention relates to methods and apparatus that use electromagnetic radiation for dermatological treatment and, more particularly to a method and apparatus that use optical radiation to ablate or damage a target area of skin surface for dermatological treatment, which skin surface includes the epidermis and parts of the dermis as the objective or side effect of the desired treatment.

**2. Background Art**

[0002] There is an increasing demand for repair of or improvement to skin defects, which can be induced by aging, sun exposure, dermatological diseases, traumatic effects, and the like. Many treatments which use electromagnetic radiation have been used to improve skin defects by

inducing a thermal injury to the skin, which results in a complex wound healing response of the skin. This leads to a biological repair of the injured skin.

[0003] Various techniques providing this objective have been introduced in recent years. The different techniques can be generally categorized in two groups of treatment modalities: ablative laser skin resurfacing ("LSR") and non-ablative collagen remodeling ("NCR"). The first group of treatment modalities, i.e., LSR, includes causing thermal damage to the epidermis and/or dermis, while the second group, i.e., NCR, is designed to spare thermal damage of the epidermis.

[0004] LSR with pulsed CO<sub>2</sub> or Er:YAG lasers, which may be referred to in the art as laser resurfacing or ablative resurfacing, is considered to be an effective treatment option for signs of photo aged skin, chronically aged skin, scars, superficial pigmented lesions, stretch marks, and superficial skin lesions. However, patients may experience major drawbacks after each LSR treatment, including edema, oozing, and burning discomfort during first fourteen (14) days after treatment. These major drawbacks can be unacceptable for many patients. A further problem with LSR procedures is that the procedures are relatively painful and therefore generally require an application of a significant amount of analgesia. While LSR of relatively small areas can be performed under local anesthesia provided by injection of an anestheticum, LSR of relatively large areas is frequently performed under general anesthesia or after nerve blockade by multiple injections of anesthetic.

[0005] Any LSR treatment results in thermal skin damage to the treatment area of the skin surface, including the epidermis and/or the dermis. LSR treatment with pulsed CO<sub>2</sub> lasers is particularly aggressive, causing thermal skin damage to the epidermis and at least to the superficial dermis. Following LSR treatment using CO<sub>2</sub> lasers, a high incidence of complications can occur, including persistent erythema, hyperpigmentation, hypopigmentation, scarring, and infection (e.g., infection with Herpes simplex virus). LSR treatment with the Er:YAG laser has been introduced as a more gentle alternative to the CO<sub>2</sub> laser, due to the lesser penetration depth of the Er:YAG pulsed laser. Using the Er:YAG laser results in a thinner zone of thermal injury within the residual tissue of the target area of the skin. However, LSR that uses the Er:YAG laser produces side effects similar to those made by LSR that uses the CO<sub>2</sub> laser within the first days after treatment.

[0006] A limitation of LSR using CO<sub>2</sub> or Er:YAG lasers is that ablative laser resurfacing generally can not be performed on the patients with dark complexions. The removal of pigmented epidermis tissue can cause severe cosmetic disfigurement to patients with a dark complexion, which may last from several weeks up to years, which is considered by most patients and physicians to be acceptable. Another limitation of LSR is that ablative resurfacing in areas other than the face generally have a greater risk of scarring. LSR procedures in areas other than the face result in an increased incidence of an unacceptable scar formation because the recovery from skin injury within these areas is not very effective.

[0007] In an attempt to overcome the problems associated with LSR procedures, a group of NCR techniques has emerged. These techniques are variously referred to in the art as non-ablative resurfacing, non-ablative subsurfacing, or non-ablative skin remodeling. NCR techniques generally utilize non-ablative lasers, flashlamps, or radio frequency current to damage dermal tissue while sparing damage to the epidermal tissue. The concept behind NCR techniques is that the thermal damage of only the dermal tissues is thought to induce wound healing which results in a biological repair and a formation of new dermal collagen. This type of wound healing can result in a decrease of photoaging related structural damage. Avoiding epidermal damage in NCR techniques decreases the severity and duration of treatment related side effects. In particular, post procedural oozing, crusting, pigmentary changes and incidence of infections due to prolonged loss of the epidermal barrier function can usually be avoided by using the NCR techniques.

[0008] Various strategies are presently applied using nonablative lasers to achieve damage to the dermis while sparing the epidermis. Nonablative lasers used in NCR procedures have a deeper dermal penetration depth as compared to ablative lasers used in LSR procedures. Wavelengths in the near infrared spectrum can be used. These wavelengths cause the non-ablative laser to have a deeper penetration depth than the very superficially-absorbed ablative Er:YAG and CO<sub>2</sub> lasers. The dermal damage is achieved by a combination of proper wavelength and superficial skin cooling, or by focusing a laser into the dermis with a high numerical aperture optic in combination with superficial skin cooling. While it has been demonstrated that these techniques can assist in avoiding epidermal damage, one of the major drawbacks of these techniques is their limited efficacies. The improvement of photoaged skin or scars after the treatment with NCR

techniques is significantly smaller than the improvements found when LSR ablative techniques are utilized. Even after multiple treatments, the clinical improvement is often far below the patient's expectations. In addition, clinical improvement is usually several months delayed after a series of treatment procedures.

[0009] Another limitation of NCR procedures relates to the breadth of acceptable treatment parameters for safe and effective treatment of dermatological disorders. The NCR procedures generally rely on an optimum coordination of laser energy and cooling parameters, which can result in an unwanted temperature profile within the skin leading to either no therapeutic effect or scar formation due to the overheating of a relatively large volume of the tissue.

[0010] Yet another problem of non-ablative procedures relates to the sparing of the epidermis. While sparing the epidermis is advantageous in order to decrease the side effects related to complete removal of the epidermis, several applications of NCR procedures may benefit from at least partial removal of epidermal structures. For example, photoinduced skin aging manifests not only by the dermal alterations, but also by epidermal alterations.

[0011] A further problem of both ablative and nonablative resurfacing is that the role of keratinocytes in the wound healing response is not capitalized upon. Keratinocyte plays an active role in the wound healing response by releasing cytokines when the keratinocyte is damaged. During traditional ablative resurfacing procedures, the keratinocytes are removed from the skin along with the epidermis, thereby removing them from the healing process altogether. On the other hand, in traditional non-ablative procedures, the keratinocytes, which are located in the epidermis, are not damaged, therefore they do not release cytokines to aid in the healing process.

[0012] Another major problem with all LSR and NCR techniques now used is the appearance of visible spots and/or edges after treatment due to inflammation, pigmentation, or texture changes, corresponding to the sites of treatment. Devices for LSR and NCR produce macroscopic (easily seen) sexposure areas. For example, laser exposure spot diameters typically vary from about 1 to 10 mm, and NCR exposure spot diameters from about 3 to 50 mm. Some devices, such as indense pulsed light devices, leave "boxes" of skin response due to rectangular output patterns on the skin. Patients do not like such spot or box patterns, easily seen as red, brown or white

areas ranging from on the order of millimeters to centimeters in size, which remain for days or even years after treatment.

[0013] Therefore, there is a need to provide a procedure and apparatus that combine safe and effective treatment for improvement of dermatological disorders with minimum side effects, such as intra procedural discomfort, post procedural discomfort, lengthy healing time, and post procedural infection.

#### SUMMARY OF THE INVENTION

[0014] It is therefore one of the objects of the present invention to provide an apparatus and method that combines safe and effective treatment for an improvement of dermatological disorders with minimum side effects. Another object of the present invention is to provide an apparatus and method that cause thermal skin damage to only a fraction of a target area of skin.

[0015] These and other objects can be achieved with the exemplary embodiment of the apparatus and method according to the present invention, in which portions of a target area to be subjected to irradiation are masked. The exemplary apparatus can include at least one shielding member configured to mask at least one portion of a target area of skin from electromagnetic radiation, in which the shielding members are formed such that a minimal amount of electromagnetic radiation is reflected back towards an electromagnetic radiation source.

[0016] In another advantageous embodiment of the present invention, electromagnetic radiation can be generated by an electromagnetic radiation source, thus causing the electromagnetic radiation to be applied to a target area of the skin. At least one portion of the target area of the skin is then masked from the electromagnetic radiation using a mask.

[0017] In yet another advantageous embodiment of the present invention, an apparatus and method for treating dermatological conditions is provided. In particular electromagnetic radiation can be generated, and a electromagnetic radiation generated by a electromagnetic radiation source may be directed (by a delivery module) to a small area within the target area of the skin, such that the small area is located relative to the delivery module. In addition, a translator can move the delivery module, such that the delivery module targets a plurality of individual exposure sites.

[0018] In a further advantageous embodiment of the present invention, the electromagnetic radiation can be applied to a first individual exposure area of the target area of the skin. The electromagnetic radiation can then be applied to a second individual exposure area of the target area of the skin.

#### BRIEF DESCRIPTION OF THE DRAWINGS

[0019] For a more complete understanding of the present invention and its advantages, reference is now made to the following description, taken in conjunction with the accompanying drawings, in which:

[0020] Figs. 1A – 1C show progressive illustrations of a first exemplary embodiment of a fractional resurfacing system for conducting various dermatological treatments at various stages of use according to the present invention;

[0021] Fig. 2 shows a top view of a first exemplary embodiment of a mask according to the present invention;

[0022] Fig. 3 shows a cross-sectional view of the mask of Fig. 2;

[0023] Fig. 4 shows a top view of a second exemplary embodiment of the mask according to the present invention;

[0024] Fig. 5 shows a cross-sectional view of the mask of Fig. 4;

[0025] Fig. 6 shows a cross-sectional view of another variant of the mask of Fig. 4;

[0026] Figs. 7A and 7B show progressive illustrations of a second exemplary embodiment of the fractional resurfacing system for conducting various dermatological treatments at various stages of use according to the present invention;

[0027] Fig. 8 shows a top view of small individual exposure areas created by the fractional resurfacing system of Figs. 7A and 7B; and

[0028] Fig. 9 shows an exemplary embodiment of a system for monitoring the location of the fractional resurfacing system of Figs. 7A and 7B.

[0029] Throughout the drawings, the same reference numerals and characters, unless otherwise stated, are used to denote like features, elements, components, or portions of the illustrated embodiments. Moreover, while the present invention will now be described in detail with reference to the Figures, it is done so in connection with the illustrative embodiments.

#### DETAILED DESCRIPTION OF THE PREFERRED EMBODIMENTS

[0030] Figs. 1A – 9 illustrate various embodiments of a method and apparatus for fractional resurfacing of a target area of skin. Generally, the exemplary methods and apparatus deliver an electromagnetic radiation to the patient's skin defined by various patterns, so as to induce thermal injury of the skin surface corresponding to such patterns and involving only a fraction of the targeted surface area of the skin. Such technique combines the efficacy of ablative resurfacing procedures with the minimal side effects of non-ablative procedures. The delivery of the electromagnetic radiation to the skin in a predetermined pattern is achieved by either masking parts of the target area of the skin surface in order to protect the masked parts of the skin surface from the electromagnetic radiation, or by utilizing a light beam of relatively small diameter which is scanned across the skin surface by various means in order to generate a specific pattern for effecting superficial thermal skin injury.

[0031] Fractional resurfacing is defined as the controlled ablation, removal, destruction, damage or stimulation of multiple small (generally less than 1 mm) individual exposure areas of skin tissue with intervening spared areas of skin tissue, performed as a treatment to improve the skin. The individual exposure areas may be oval, circular, arced and/or linear in shape. The spatial scale of fractional resurfacing is chosen to avoid the appearance of various spots or boxes on a macroscopic scale, while still providing effective treatment because the multiple small areas can be exposed to greater than a minimal stimulus. For example, removal or photothermal destruction of thousands of 0.1 mm diameter individual exposure areas, spaced 0.2 mm apart, and extending into the skin up to a depth of 0.5 mm, is well tolerated and produces effective improvement of photoaging, without apparent spots and with rapid healing. Spared skin between the individual exposure areas rapidly initiates a wound healing response, which is better tolerated than conventional LSR.

[0032] During the exemplary fractional resurfacing procedure of the present invention, certain portions of the target area remain undamaged, thereby preserving keratinocytes and melanocytes, which serve as a pool of undamaged cells to promote reepithelialization. This procedure differs from the traditional resurfacing procedures, such that the entirety of the target area is damaged. In traditional resurfacing procedures, reepithelialization is generally initiated from the depth of an undamaged follicular epithelium. Because the traditional procedures remove the entire epithelium, an important factor for the time of reepithelialization is the density of follicles. The vellus hair density of the face ( $439$  hairs/cm $^2$ ) of the subject is significantly higher than on the back of the subject ( $85$  hairs/cm $^2$ ). Therefore, the face of the subject, generally experiences better and faster reepithelialization in comparison to other body areas with a lower hair density.

[0033] The resurfacing of the dark pigmented skin is currently not very frequently performed because of the prolonged repigmentation process. The fractional resurfacing technique improves the repigmentation process but, melanocytes do not migrate well. By sparing certain portions of the target area of the skin, the travel distance of melanocytes can be decreased, thereby reducing the repigmentation time and allowing the resurfacing of all skin types.

[0034] Figs. 1A – 1C illustrate a progressive use of a first exemplary embodiment of a fractional resurfacing system 100 for conducting various dermatological treatments using electromagnetic radiation (“EMR”) and generating a superficial pattern of skin damage of a target area by using a mask according to the present invention. The system 100 may be used for collagen remodeling, removal of unwanted pigment or tattoo, and/or other dermatological applications. As shown in Figs. 1A-1C, the system 100 includes a case 101, a control module 102, an EMR source 104, delivery optics 106 and a mask 108. The case 101 contains the control module 102, the EMR source 104, and the delivery optics 106. An aperture is provided through a sidewall of the case 101. The mask 108 is placed in registration with the aperture formed through the sidewall of the case 101. By placing the mask 108 in registration with the aperture of the case 101, the focal length of the EMR emitted by the delivery optics 106 is fixed, and can be configured such that it does not impact the side of the mask 108, so as to cause injuries to the operator of the fractional ablation system 100. The control module 102 is in communication with the EMR source 104, which in turn is operatively connected to the delivery optics 106.

**[0035]** In one exemplary variant of the present invention, the control module 102 can be in wireless communication with the EMR source 104. In another variant, the control module 102 may be in wired communication with the EMR source 104. In another exemplary variant of the present invention, the control module 102 can be located outside of the case 101. In another variant, the EMR source 104 is located outside of the case 101. In still another variant, the control module 102 and the EMR source 104 are located outside of the case 101. It is also possible that the mask 108 is not connected to the case 101.

**[0036]** The control module 102 provides application specific settings to the EMR source 104. The EMR source 104 receives these settings, and generates EMR based on these settings. The settings can control the wavelength of the EMR, the energy delivered to the skin, the power delivered to the skin, the pulse duration for each EMR pulse, the fluence of the EMR delivered to the skin, the number of EMR pulses, the delay between individual EMR pulses, the beam profile of the EMR, and the size of the area within the mask exposed to EMR. The energy produced by the EMR source 104 can be an optical radiation, which is focused, collimated and/or directed by the delivery optics 106 to the mask 108. The mask 108 can be placed on a target area of a patient's skin, and may provide a damage pattern on the target area of the skin with a fill factor in the range from 0.1% to 90%. The fill factor is the percentage of the target area exposed to the EMR that is emitted by the EMR source 106.

**[0037]** In one exemplary embodiment, the EMR source 106 is one of a laser, a flashlamp, a tungsten lamp, a diode, a diode array, and the like. In another exemplary embodiment, the EMR source 106 is one of a CO<sub>2</sub> laser and a Er:YAG laser.

**[0038]** Prior to being used in a dermatological treatment, the system 100 shown in Fig. 1A can be configured by a user. For example, the user may interface with the control module 102 in order to specify the specific settings usable for a particular procedure. The user may specify the wavelength of the EMR, the energy delivered to the skin, the power delivered to the skin, the pulse duration for each EMR pulse, the fluence of the EMR delivered to the skin, the number of EMR pulses, the delay between individual EMR pulses, the beam profile of the EMR, and the size of the area within the mask exposed to EMR. The EMR source 104 may be set to produce a collimated pulsed EMR irradiation with a wavelength ranging from 400 to 11,000 nm, and

preferably near 3.0  $\mu\text{m}$  when using an Er:YAG laser and near 10.6  $\mu\text{m}$  when using a CO<sub>2</sub> laser as the EMR source. The collimated pulsed EMR irradiation may be applied which has a pulse duration in the range of 1  $\mu\text{s}$  to 10 s, preferably in the range of 100  $\mu\text{s}$  to 100 ms, and more preferably in the range of 0.1 ms to 10 ms, and fluence in the range from 0.01 to 100 J/cm<sup>2</sup>, and preferably in the range from 1 to 10 J/cm<sup>2</sup>. The applied EMR should be able to achieve at least a temperature rise within the exposed areas of the skin that is sufficient to cause thermal damage to the epidermis 110 and/or the dermis 112. The peak temperature sufficient to cause thermal damage in the exposed tissues is time dependant and at least in the range of 45° C to 100° C. For exposure times in the range of 0.1 ms to 10 ms the minimum temperature rise required to cause thermal damage is in the range of approximately 60° C to 100° C. The depth of thermal damage can be adjusted by proper choice of wavelength, fluence per pulse and number of pulses.

[0039] During the dermatological treatment, the system 100 produces EMR 120 which is directed to the target area of the skin 114, as shown in Fig. 1B. The EMR 120 may be pulsed multiple times to create the appropriate effect and irradiation in the target area of the skin 114.

[0040] After the dermatological treatment is completed, the target area of the skin 114 is likely damaged in specific places. The application of the EMR 120 creates a prearranged thermal skin damage 130 in an epidermal tissue 110 and the dermal tissue 112. The mask 108 controls in a location where the thermal skin damage 130 is created. The thermal skin damage 130 generally accounts for only 0.1% to 90% of the skin surface area in the target area. A fill factor is defined as the ratio of surface area of the target area of skin thermally damaged by EMR to surface area of the target area of the skin.

[0041] Fig. 2 illustrates a top view of a first exemplary embodiment of the mask 108 according to the present invention. The mask 108 includes shielding structured 202. The diameter of the mask 108 should preferably be matched to greater than the size of the diameter of the target area. The target area is defined as the area targeted by the collimated EMR emitted by the EMR source 104, which can be in the range 1 - 100 mm in diameter, preferably within the range of 5 to 20 mm. This diameter of most of the currently commercially available CO<sub>2</sub> and Er:YAG laser systems can match the diameter of the exposed area. The width of shielding structures 202 within the mask 108 should be in the range of 50 to 300  $\mu\text{m}$ . The width of the apertures of the

mask 108 that are formed by the shielding structures should be in the range of 10 - 1000  $\mu\text{m}$ , and preferably in the range of 50 to 300  $\mu\text{m}$ . The shielding-exposure ratio surface area covered by the shielding structures 202 to the surface area exposed by the apertures effects the clinical efficacy and provides side effects of the dermatological treatment. This also determines the fill factor and the pattern of the thermal damage of the skin. The depth of thermal damage is determined by the number of pulses, the fluence of the EMR and the wavelength of the EMR. The shielding-exposure ratio of the mask 108 will vary for different dermatological treatments, particular patient needs, particular patient indications, skin types and body areas.

[0042] The mask 108 may have a large shielding-exposure ratio at the edge of the mask 108 to generate a transition zone at the edge of resurfaced area. This technique is called "feathering." It avoids a sharp macroscopically visible demarcation between treated and untreated areas. In another preferred embodiment, a mask may be used that has a large shielding-exposure ratio at the edge of a conventionally resurfaced area to generate a transition zone.

[0043] The surface of the mask 108 should preferably have a minimal absorption at the wavelength generated by the EMR source 104 for the particular dermatological process. Such absorption can decrease the undesirable heating of the mask 108. The mask 108 may be coated by a metal material for effectuating a minimal absorption of the EMR. The design of the shielding structures 202 of the mask 108, a cross-section A-A of which is shown in Fig. 3, generally takes into consideration safety aspects, including a back-reflected EMR in order to avoid EMR inflicted accidents. The shielding structures 202 are shaped in a peaked manner to minimize the amount of back reflected EMR. Also, with the mask 108 being connected to the case 101 the distance between the delivery optics 106 and the mask 108 is fixed, thereby minimizing the chances that EMR would be reflected back towards the user by hitting the edge of the mask 108. Additionally, the microstructure of the mask 108 can have a periodicity preferably in the range of the wavelength of the EMR emitted by the delivery optics 106. This configuration can diffuse the collimated EMR emitted by the delivery optics 106 into a highly scattered beam so as to decrease the risk of EMR-related accidents.

[0044] In one exemplary embodiment, the metal coating of the mask 108 may be composed of gold, silver, or copper materials, or the like. In another exemplary embodiment, the

microstructure of the surface of the mask 108 may have a periodicity in the range of the wavelength of the EMR emitted by the delivery optics 106.

[0045] The mask 108 may also have a configuration so as to provide effective skin cooling during the exposure thereof with the EMR radiation. Skin cooling provides significant anesthetic effects, and has other advantages related to the pattern induced by the EMR radiation. The mask 108 can be cooled prior to the beginning of the dermatological procedure, during the procedure by spraying an evaporative agent or a precooled liquid onto the mask 108 between the successive EMR pulses, or during the procedure by introducing a cool or cold liquid into microchannels 302 (shown in Fig. 3) running through the mask 108. The cooling of the mask 108 has a secondary advantage in that such cooling of the mask 108 decreases the rate of the EMR absorption by the mask 108, as the rate of the EMR absorption by the metals increases with the increasing temperature.

[0046] In order to provide skin cooling as described above, the temperature of the mask 108 should be in the range of 37° C to -20° C, and preferably 10° C to -4° C. The mask 108 can both protect and cool the portions of the skin surface that are not exposed to EMR emitted by the EMR source 104. In addition to cooling and shielding portions of the skin surface, the mask 108 allows the debris ejected during ablative procedures to escape, and thereby not interfere with the beam delivery for successive pulses.

[0047] Fig. 3 illustrates a cross-section A-A of the mask 108 of Fig. 2. The cross-section A-A shows the microchannels 302 that run through at least the shielding structures 202 of the mask 108. A cooling agent, e.g., either a liquid or gas, may circulate through these microchannels 302 during a dermatological procedure, thereby removing heat from the protected skin and the mask 108 itself.

[0048] Fig. 4 illustrates a top view of a second embodiment of the mask 400 according to the present invention. The mask 400 differs from the mask 108 only in the layout and design of the shielding structures 402. The details of the mask 400 are in all other respects substantially similar to those of the mask 108. The shielding structures 402 are cylindrical in shape, as indicated in cut-away cross-sections B-B and C-C, shown in Figs. 5 and 6, respectively. The shielding structures 402 of the mask 400 contain microchannels 502 and 602, which are capable

of carrying a cooled liquid or gas so as to cool the mask 400 and the masked portions of the target area of the skin. The microchannels 502, 602 intersect at the intersection of the shielding structures 402.

[0049] In an exemplary embodiment of the present invention, the microstructures 502, 602 are not required to intersect at the intersection of the shielding structures 402.

[0050] In an exemplary embodiment of the present invention, the mask 108 is an ablative mask. An ablative mask includes multiple sections having various thicknesses. Prior to a procedure, the ablative mask is attached to the skin with an adhesive. During the procedure having multiple EMR pulses, the ablative mask is ablated, such that the thickness of each of the multiple sections is diminished, potentially gradually exposing different areas of the skin to the EMR pulses. The ablative mask can be composed of various materials including polymer materials. The ablative mask can be easily produced by imprinting a pattern therein.

[0051] A particular dermatological treatment, i.e., the removal of tattoos, shall be described in further detail. Tattoo removal may be performed with a combination of an ablative EMR and the mask 108. In particular, utilizing the CO<sub>2</sub> laser and/or the Er:YAG laser may be appropriate for this application. During this dermatological procedure, the tattoo can be exposed to ablative EMR radiation with the mask 108 providing a fill factor of the target area in the range of 10 to 90%, and preferably in the range of 25 to 70%. Preferably, the mask 108 is applied under pressure to the skin, which minimizes the blood flow during the procedure. Limiting the blood flow during the procedure allows a deeper ablation of the skin surface before blood can interfere with the EMR radiation, thereby limiting the ablation depth. Multiple pulses of ablative EMR radiation can be applied to the individual areas of the tattoo until the desired ablation depth is reached. The desired ablation depth can be in the range of 100  $\mu$ m to 5 mm. This exemplary procedure can cause a specific fraction of the tattoo that is controlled by the mask 108 to be immediately ablated. Wound healing may be enhanced because only a fraction of the surface is ablated.

[0052] The removal of tattoos utilizing fractional resurfacing may be augmented using a short pulsed EMR, preferentially absorbed by the tattoo particles either before or after the application of the fractional resurfacing. In a short pulsed-laser application, the laser may be pulsed for short

periods of time, preferably for less than 1  $\mu$ s in duration. The EMR source used in this type of procedure can preferably be a Q-switched ruby laser, a Nd:YAG laser, a KTP laser and/or an Alexandrite laser. The objective of this procedure is to release the pigment within areas that are not exposed to fractional resurfacing ablation. The released pigment particles may drain in the ablated channels, and can be flushed from the area after the procedure by the blood resident in the target area and/or an external rinsing agent, e.g., saline. Several such procedures may be utilized until the desired clearance of the tattoo has occurred.

[0053] As an alternative to the fractional resurfacing using a mask, a second embodiment of a fractional resurfacing system 700, as shown as the progressive use thereof in Figs. 7A – 7B, can be used. The system 700 can include a case 701, a control module 702, an electromagnetic radiation ("EMR") source 704, delivery optics 706, an x-y translator 708 and an optically transparent plate 709. The case 701 may contain the control module 702, the EMR source 704, the delivery optics 706 and the translator 708. As with the system 100, an aperture may be formed through a sidewall of the case 701. The optically transparent plate 709 may be placed in registration with the aperture that is formed through the sidewall of the case 701. Placing the plate 709 in registration with the aperture formed through the sidewall of the case 701 seals the system 700, which contains sophisticated translation mechanisms, e.g., the delivery optics 706 and the translator 708. The control module 702 is in communication with the translator 708 and the EMR source 704, and the EMR source 704 is operatively connected to the delivery optics 706.

[0054] In one exemplary variant of the present invention, the control module 702 can be located outside of the case 701. In another exemplary variant, the EMR source 704 is located outside of the case 701. In still another variant, the control module 702 and the EMR source 704 are located outside of the case 701.

[0055] The control module 702 provides application specific settings to the EMR source 704, and controls the x-y translator 708. The EMR source 704 receives these settings, and generates EMR based on these settings. The settings can control the wavelength of the energy produced, the intensity of the energy produced, the fluence of the energy produced, the duration of the dermatological procedure, the pulse length of each of the EMR pulses administered during the

procedure, the spatial distance between individual exposure areas 716 (shown in Fig. 8), the shape of individual exposure areas 716, the pattern defined by individual exposure areas 716, and the fill factor of the target area. The EMR source 704 can be a laser or other light source. The EMR produced by the EMR source 704 can be delivered through a fiber, waveguide or mirrors if the source is located outside the delivery optics 706. Alternatively, if the EMR source 704 is located in a close vicinity to the skin 714, the EMR source 704 produces the EMR directly to the delivery optics 706. The energy produced by the EMR source 704 may be focused and/or directed by focusing optics in the delivery optics 706 to one of the an individual exposure areas 716, shown in Fig. 8. Each of the individual exposure areas 716 are located within the target area of the skin 714, and are relatively small compared to the target area of the skin 714. The target area of the skin 714 can generally be 1 cm<sup>2</sup> in size.

[0056] In one embodiment of the present invention, the focusing optics of the delivery optics 706 may contain a beam collimator.

[0057] Prior to use in a dermatological treatment and similarly to the use of system 100, the system 700, as shown in Fig. 7A, can be configured by a user. In particular, the user interfaces with the control module 702 in order to specify the specific settings to be used for a particular procedure. The user may specify the desired damage pattern, the wavelength of the energy produced by the EMR source 704, the intensity of the energy produced, the fluence of the energy produced, the length of time the treatment will take and the pulse duration of the EMR source 704. During the treatment, the translator 708 moves the delivery optics 706 across sequential portions of the target area of the skin 714 in order to treat the entire target area. When one of the portions of the target area has been completely treated, the system 700 is moved to the next portion of the target area. For example, the system 700 is moved at the completion of irradiation of each portion of the target area until the desired skin surface damage pattern is achieved for the entire area. The system 700 can be moved using discrete movements from one sequential portion to the next, i.e., stamping mode, or using continuous movement across the skin surface, i.e., continuous scanning mode. In either case, the movement of the delivery optics 706, driven by the translator 708, is controlled by the control unit 702 and likely matched with the movement of the system 700 by the operator (or the user) in order to provide the desired surface damage pattern to the target area of the skin 714.

[0058] In a preferred embodiment, the system 700 includes a position sensor, which is in communication with the control module 702. The position sensor is capable of sensing the relative velocity as between the skin 114 and the case 701. The position sensor can be an optical mouse, wheels, track ball, conventional mouse, and the like.

[0059] Before the initiation of a dermatological procedure, the optically transparent plate 709 can be brought in a direct contact with the skin surface covering the target area. The optically transparent plate 709 can be composed out of any material having good thermal conductivity, and being transparent over a broad range of the visible and near infrared spectrum. The plate 709 seals the system 700, which contains sophisticated translation mechanisms, and provides cooling to the target area of the skin 714. The plate 709 can provide cooling to the target area of the skin 714 in two ways: heat conduction and heat convection. Heat conduction transfers heat through the optically transparent plate 709 to the case 701, which provides cooling by circulating a coolant agent through the case 701 of the system 700. The entire optically transparent plate 709 can also be cooled prior to application to the target area of the skin 714. Alternatively, heat convection can be utilized for this procedure. An evaporating agent sprayed onto the optical window or onto a compartment in good thermal contact with the window may also be utilized. The delivery of the evaporating agent can be administered during the procedure between EMR pulses through a valve, which can be controlled by a thermostat with a temperature sensor at the optical plate.

[0060] In one embodiment, of the present invention the optically transparent plate 709 can be composed of sapphire or quartz. In another embodiment of the present invention, the system 700 can be moved multiple times over the same portion of the skin 714 until the desired fill factor is achieved. In yet another embodiment, multiple procedures can be performed to achieve the desired effect.

[0061] During the dermatological procedure, the EMR source 704 emits EMR having a wavelength in the range of 400 - 12,000 nm. Preferably the EMR has a wavelength in one of the following ranges: 1,300 to 1,600 nm, 1,850 to 2,100 nm, 2,300 to 3,100 nm and around 10,640 nm. Depending on the application, a single wavelength or a combination of different wavelengths may be utilized. The EMR source 704 can be a diode laser, a fiber laser, a solid

state laser, a gas laser, and the like. The pulse duration can range from 100  $\mu$ s to 100 ms, and preferably in the range from 500  $\mu$ s to 15 ms. The energy density per pulse within an individual exposure area 716 may be in the range of 0.1 to 100 J/cm<sup>2</sup>, and preferably 1 to 20 J/cm<sup>2</sup>.

[0062] After the dermatological treatment is completed, the target area of the skin 714 is damaged in a specific pattern. The application of EMR creates the thermal skin damage in an epidermis 710 and a dermis 712 of the skin 714. The radiation provided by the EMR source 704 is delivered to the skin 714 within multiple small individual exposure areas 716, shown in Fig. 7B, through the delivery optics 706. The delivery optics 706 can deliver multiple individual beams across the target area of the skin surface.

[0063] Fig. 8 illustrates a top view of the small individual exposure areas 716 of the epidermis. The shape of the individual exposure areas 716 may be circular (shown in Fig. 8), elliptical, rectangular, linear or irregular with a lateral diameter of the smallest dimension in the range of 1 – 500  $\mu$ m. The fill factor of the target area can be approximately 20 – 40%.

[0064] The system 700 can create multiple individual exposure areas 716 through ablation, removal, photothermal coagulation, thermal necrosis and/or stimulation. The multiple areas can be exposed sequentially or simultaneously. Sequential exposure may be achieved by scanning or moving an energy source which may be either pulsed, shuttered or continuous. Simultaneous exposure can be achieved, for example, by an array of sources or a multi-array of lenses. The array of sources may be a uni-dimensional array, a bi-dimensional array or the like. The array can be moved relative to the skin, and one or multiple passes of treatment can be performed in a target area.

[0065] Fig. 9 illustrates an exemplary embodiment of a monitoring system 900 according to the present invention. The monitoring system 900 tracks the movement of the system 700, and feeds such positional information to the control module 702. The control module 702 utilizes this information to appropriately instruct the translator 708 to position the delivery optics 706, such that the appropriate damage pattern is achieved across the target area of the skin 714. The monitoring system 900 may use a computer 902, a mouse 904, and a charge coupled device ("CCD") camera 906. In particular, the computer 902 receives the positional information about the system 700 from the CCD camera 906. The computer then updates the control module 702

based on this positional information as to the current position of the system 700. The control module 702 utilizes this information to cause the system 700 to create the appropriate damage pattern on the skin 714 within the target area. The mouse 904 can be an optical mouse. In addition, the monitoring system can utilize additional motion detecting devices, including, wheels or any other motion sensor.

[0066] The shape of the individual exposure areas 716 and the relative pattern represented by all of the individual exposure areas 716 may vary. The individual exposure areas 716 can have a circular, elliptical, rectangular, linear or irregular shape. The average distance between individual regions of unexposed skin surface may be in the range between 10 to 2000  $\mu\text{m}$ , and preferably in the range of 100 to 500  $\mu\text{m}$ . The macroscopic pattern of the individual exposure areas 716 may be a field of uniformly distributed individual exposure areas 716 with constant spacing throughout the target area, randomly distributed individual exposure areas 716 within the target area, and/or regularly distributed individual exposure areas 716 with constant average spacing with randomly shifted location. In particular, having regularly distributed individual exposure areas 716 with constant average spacing with randomly shifted location may be useful to minimize undesirable effects, which may occur during multiple treatments. Such multiple treatments are utilized to cover the entire area as homogeneously as possible by the individual exposure areas 716 during the course of multiple treatments. However, uniformly distributed individual exposure areas 716 with constant spacing throughout the target area may create unwanted spatial distributions similar to moiré patterns, resulting in spatial interference macroscopic patterns generated with a distance in between the areas of exposure which have a significant spatial period. In order to minimize the occurrence of moiré patterns, a randomized shift within the range of 10 to 50% of the average distance between individual exposure areas 716 during a single scan may be utilized.

[0067] The treatment can be performed in by a single treatment covering the skin surface with a specific surface damage pattern, or by multiple treatments either performed at the same visit or during different treatment visits. Individual or multiple exposures can be used to achieve the appropriate thermal damage in particular individual exposure areas 716.

[0068] Fractional resurfacing may cause portions of the epidermis to be thermally damaged or ablated, thereby reducing the efficacy of the barrier function of the epidermis and in particular decreasing the stratum corneum. This facilitates the delivery of drugs or specific substances to the dermis and epidermis which can either enhance the effects of the treatment, or decrease the side effects caused by partial damage of the epidermis and/or dermis. Groups of drugs and substances, which may enhance the efficacy of skin remodeling include growth factors, collagen byproducts, collagen precursors, hyaluronic acid, vitamins, antioxidants, amino acids and supplemental minerals among others. Groups of drugs and substances, which may decrease side effects, can be steroidal anti-inflammatory drugs, non-steroidal anti-inflammatory drugs, antioxidants, antibiotics, antiviral drugs, antiyeast drugs and antifungal drugs.

[0069] In an exemplary embodiment of the present invention, the vitamins that are used may be vitamin C and/or vitamin E. The supplemental minerals used are copper and zinc. The antioxidants can be vitamin C and/or vitamin E.

[0070] In a clinical observation, enhanced wound healing was observed for fractional resurfacing as compared to conventional resurfacing. The forearm skin of a white, male Caucasian was exposed to pulsed CO<sub>2</sub> laser radiation with identical settings of the illuminating laser beam, a Coherent Ultra Pulse Laser, CPG handpiece, at approximately 300mJ/pulse. One area was exposed to the laser beam without benefit of a mask while another area was partially shielded by a cooled mask. More pronounced erythema was evident at the conventionally resurfaced test site as compared to the fractionally resurfaced test site.

[0071] The fill factor of the target area may be monitored by sensing the electrical impedance of the skin from a location on the skin within the target area to a remote location on the skin outside of the target area during or after treatment. An indicator capable of staining the defects in the stratum corneum (for example, trypan glue) or transdermal waterloss are effective indicators of the fill factor of the target area.

[0072] The foregoing merely illustrates the principles of the invention. Various modifications and alterations to the described embodiments will be apparent to those skilled in the art in view of the teachings herein. It will thus be appreciated that those skilled in the art will be able to

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devise numerous techniques which, although not explicitly described herein, embody the principles of the invention and are thus within the spirit and scope of the invention.

WHAT IS CLAIMED IS:

1. An apparatus, comprising:  
at least one member configured to mask at least one portion of a target area of skin from an electromagnetic radiation provided by an electromagnetic radiation source, wherein at least one member is configured such that a particular amount of the electromagnetic radiation that impacts the at least one member is reflected in a direction of the electromagnetic radiation source.
2. The apparatus of claim 1, wherein the target area is a predetermined area of the skin.
3. The apparatus of claim 1, wherein each of the at least one shielding member is configured to reflect the impacted electromagnetic radiation away from the apparatus.
4. The apparatus of claim 1, wherein each of the at least one shielding member is configured to absorb a minimal amount of electromagnetic radiation.
5. The apparatus of claim 1, wherein the electromagnetic radiation is optical radiation.
6. The apparatus of claim 1, wherein the electromagnetic radiation source is an ablative laser.
7. The apparatus of claim 1, wherein the electromagnetic radiation source is generated by a carbon dioxide laser.
8. The filtering apparatus of claim 1, wherein the electromagnetic radiation source is an Er:YAG laser.
9. The apparatus of claim 1, wherein the at least one shielding member masks at least 0.1% of the target area from the electromagnetic radiation.
10. The apparatus of claim 1, wherein the at least one member masks at most 90% of the target area from the electromagnetic radiation.
11. The apparatus of claim 1, wherein the at least one member masks the at least one portion of the target area such that the electromagnetic radiation is prevented from effecting the at least one portion of the target area.

12. The apparatus of claim 1, wherein the at least one member is at least 50  $\mu\text{m}$  in width and at most 300  $\mu\text{m}$ .
13. The apparatus of claim 1, wherein the at least one member is configured to define at least one aperture.
14. The apparatus of claim 13, wherein the at least one aperture has a width of at least 50  $\mu\text{m}$  and at most 1000  $\mu\text{m}$ .
15. The apparatus of claim 1, wherein the at least one member is cooled.
16. The apparatus of claim 1, wherein the at least one member is adapted to be cooled to at least 37°C and at most negative 20°C .
17. The apparatus of claim 1, wherein the at least one member includes at least one channel extending therethrough.
18. The apparatus of claim 17, wherein the at least one channel is configured to facilitate a cooling agent.
19. A method for treating dermatological conditions, comprising:  
controlling an electromagnetic radiation source to generate an electromagnetic radiation;  
causing the electromagnetic radiation to be applied to a target area of skin; and  
masking at least one portion of the target area of the skin from the electromagnetic radiation.
20. The method of claim 19, wherein the masking step is performed using a mask which includes at least one member.
21. The method of claim 20, wherein the at least one shielding member masks at least 0.1% of the target area from the electromagnetic radiation.
22. The method of claim 20, wherein the at least one member masks at most 90% of the target area from the electromagnetic radiation.
23. The method of claim 20, wherein the at least one member masks the at least one portion

of the target area such that the electromagnetic radiation is prevented from effecting the at least one portion of the target area.

24. The method of claim 20, wherein the at least one member masks the at least one portion of the target area such that the electromagnetic radiation has an effect on the at least one portion than an effect to other portions of the target area.

25. The method of claim 20, wherein the at least one member is at least 50  $\mu\text{m}$  in width and at most 300  $\mu\text{m}$ .

26. The method of claim 20, wherein the at least one member is configured to define at least one aperture.

27. The method of claim 26, wherein the at least one aperture has a width of at least 50  $\mu\text{m}$  and at most 1000  $\mu\text{m}$ .

28. The method of claim 20, wherein the at least one member is cooled.

29. The method of claim 20, wherein the at least one member is adapted to be cooled to at least 37°C and at most negative 20°C .

30. The method of claim 20, wherein the at least one member includes at least one channel extending therethrough.

31. The method of claim 30, wherein the at least one channel is configured to facilitate a cooling agent.

32. The method of claim 19, wherein the mask is configured to reflect a predetermined amount of the electromagnetic radiation in a direction of the electromagnetic radiation source.

33. The method of claim 19, wherein the mask is configured to reflect the electromagnetic radiation away from the electromagnetic radiation source.

34. The method of claim 19, wherein the mask is configured to diffuse the electromagnetic radiation.

35. The method of claim 19, wherein the electromagnetic radiation has a particular wavelength.
36. The method of claim 35, wherein a surface of the mask has a microstructure having a periodicity approximately in the range of the particular wavelength.
37. The method of claim 19, wherein the mask is configured to absorb a predetermined amount of the electromagnetic radiation.
38. The method of claim 19, wherein the electromagnetic radiation source is an ablative laser.
39. The method of claim 19, wherein the electromagnetic radiation source is a carbon dioxide laser.
40. The method of claim 19, wherein the electromagnetic radiation source is a Er:YAG laser.
41. The method of claim 19, further comprising the steps of:  
controlling a further electromagnetic radiation source to generate a further electromagnetic radiation; and  
applying the further electromagnetic radiation to the target area of the skin.
42. The method of claim 41, wherein the further electromagnetic radiation source is substantially the same as the electromagnetic radiation source.
43. The method of claim 41, wherein the further electromagnetic radiation source is different than the electromagnetic radiation source.
44. The method of claim 41, wherein the further electromagnetic radiation source is one of a Q-switched ruby laser, a Nd:YAG laser, a KTP laser and an Alexandrite laser.
45. The method of claim 19, further comprising the step of introducing a substance to the target area, wherein the substance is one of growth factors, collagen byproducts, collagen precursors, hyaluronic acid, vitamins, antioxidants, amino acids and supplemental minerals.
46. An apparatus for treating dermatological conditions, comprising:

a delivery module configured to direct an electromagnetic radiation generated by an electromagnetic radiation source to a target area of skin; and

a mask including at least one member configured to mask at least one portion of the target area of the skin from the electromagnetic radiation.

47. The apparatus of claim 46, wherein the at least one member is configured to reflect a predetermined amount of the electromagnetic radiation in the direction of the electromagnetic radiation source.

48. The apparatus of claim 46, wherein each of the at least one member is configured to reflect the electromagnetic radiation away from the electromagnetic radiation source.

49. The apparatus of claim 46, wherein each of the at least one member is configured to diffuse the electromagnetic radiation.

50. The apparatus of claim 46, wherein the electromagnetic radiation has a particular wavelength.

51. The apparatus of claim 50, wherein each of the at least one member includes a microstructure having a periodicity in the range of the particular wavelength.

52. The apparatus of claim 46, wherein each of the at least one member is configured to absorb a minimal amount of the electromagnetic radiation.

53. The apparatus of claim 46, wherein the electromagnetic radiation source is an ablative laser.

54. The apparatus of claim 46, wherein the electromagnetic radiation source is a carbon dioxide laser.

55. The apparatus of claim 46, wherein the electromagnetic radiation source is a Er:YAG laser.

56. The apparatus of claim 46, wherein the at least one shielding member masks at least 0.1% of the target area from the electromagnetic radiation.

57. The apparatus of claim 46, wherein the at least one member masks at most 90% of the target area from the electromagnetic radiation.

58. The apparatus of claim 46, wherein the at least one member masks the at least one portion of the target area such that the electromagnetic radiation is prevented from effecting the at least one portion of the target area.

59. The apparatus of claim 46, wherein the at least one member masks the at least one portion of the target area such that the electromagnetic radiation is prevented from effecting the at least one portion of the target area.

60. The apparatus of claim 46, further comprising a case having an aperture formed in a sidewall of the case, wherein the case contains the electromagnetic radiation source and the delivery module, and wherein the at least one member is in registration with the apertured.

61. The apparatus of claim 46, wherein the delivery module includes a beam collimator.

62. The apparatus of claim 46, wherein the delivery module includes optical components.

63. An apparatus for treating dermatological conditions, comprising:  
a delivery module configured to direct an electromagnetic radiation generated by an electromagnetic radiation source to a predetermined area within a target area of skin, wherein the predetermined area is located in a location relative to the delivery module; and  
a translator capable of moving the delivery module, such that the delivery module targets a plurality of individual exposure sites of the predetermined area.

64. The apparatus of claim 63, wherein the electromagnetic radiation source is an ablative laser.

65. The apparatus of claim 63, wherein the electromagnetic radiation source is one of a diode laser, a fiber laser, a solid state laser and a gas laser.

66. The apparatus of claim 63, further comprising a case having an aperture formed in a sidewall of the case, wherein the case contains the electromagnetic radiation source, the delivery module and the translator.

67. The apparatus of claim 66, further comprising a transparent plate in registration with the aperture, wherein the transparent plate seals the case.

68. The apparatus of claim 67, wherein the electromagnetic radiation has a particular wavelength.

69. The apparatus of claim 68, wherein the transparent plate absorbs a predetermined amount of the electromagnetic radiation at the particular wavelength.

70. The apparatus of claim 67, wherein the transparent plate is cooled to provide an aesthetic effect to the target area of the skin.

71. The apparatus of claim 67, wherein the transparent plate is configured to be cooled to at least 37°C and at most negative 20°C.

72. The apparatus of claim 63, wherein the delivery module includes a beam collimator.

73. The apparatus of claim 63, wherein the delivery module includes optical components.

74. A method for treating dermatological conditions, comprising:  
controlling an electromagnetic radiation source to generate an electromagnetic radiation;  
causing the electromagnetic radiation to be applied to a first individual exposure area of a target area of skin; and  
causing the electromagnetic radiation to be applied to a second individual exposure area of the target area of the skin.

75. The method of claim 74, wherein the first individual exposure area has a width of at least 1 μm in width and at most 500 μm.

76. The method of claim 74, wherein the target area has a surface area of approximately 1 cm<sup>2</sup>.

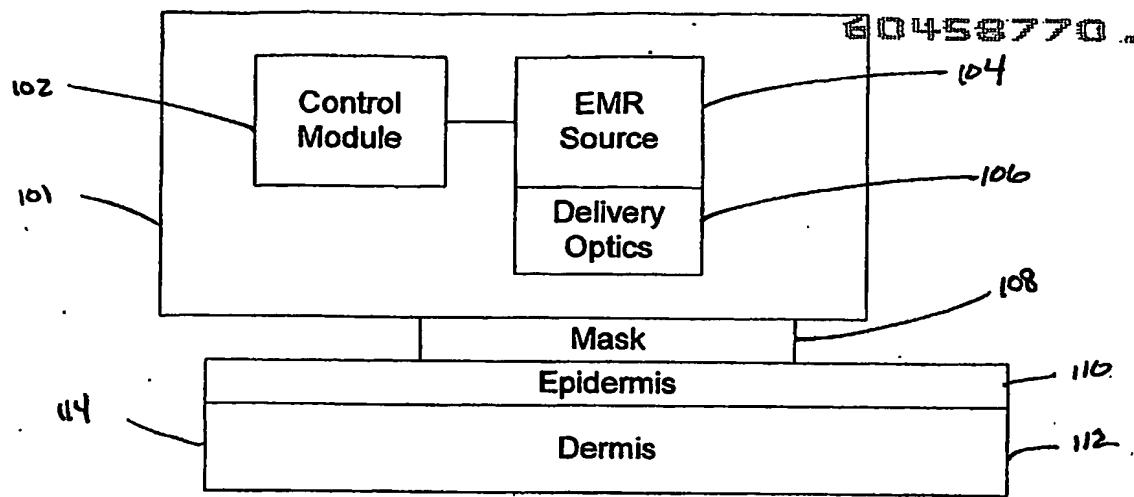
77. The apparatus of claim 74, wherein the electromagnetic radiation source is an ablative laser.

78. The apparatus of claim 74, wherein the electromagnetic radiation source is one of a diode laser, a fiber laser, a solid state laser and a gas laser.

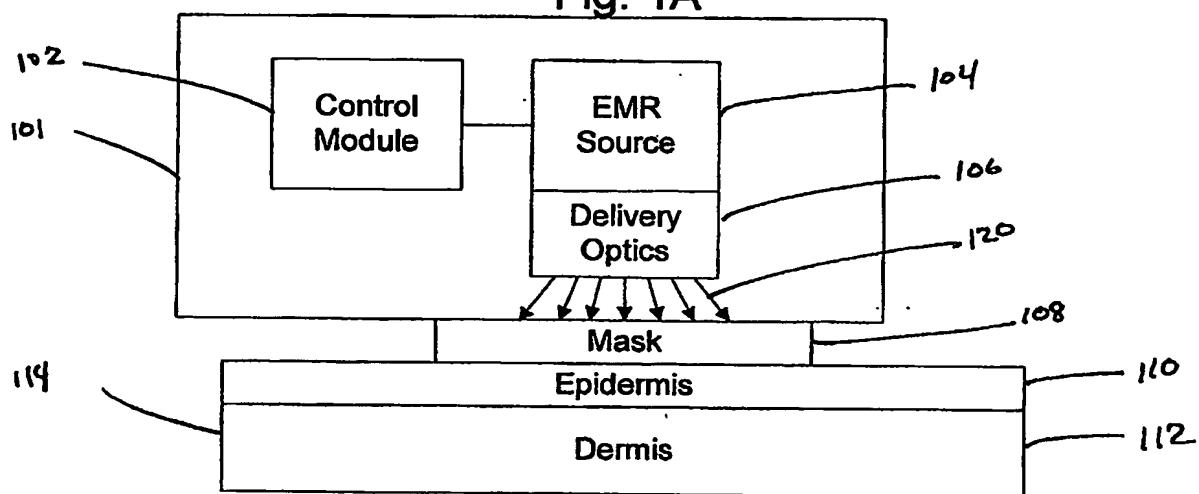
ABSTRACT

[0073] A system and method for performing fractional resurfacing of a target area of skin using electromagnetic radiation are provided. An electromagnetic radiation is generated by an electromagnetic radiation source. The electromagnetic radiation is caused to be applied to a particular portion of a target area of skin. The electromagnetic radiation can be impeded from effecting another portion of the target area of the skin by a mask. Alternatively, the electromagnetic radiation may be applied to portions of the target area of the skin, other than the particular portion.

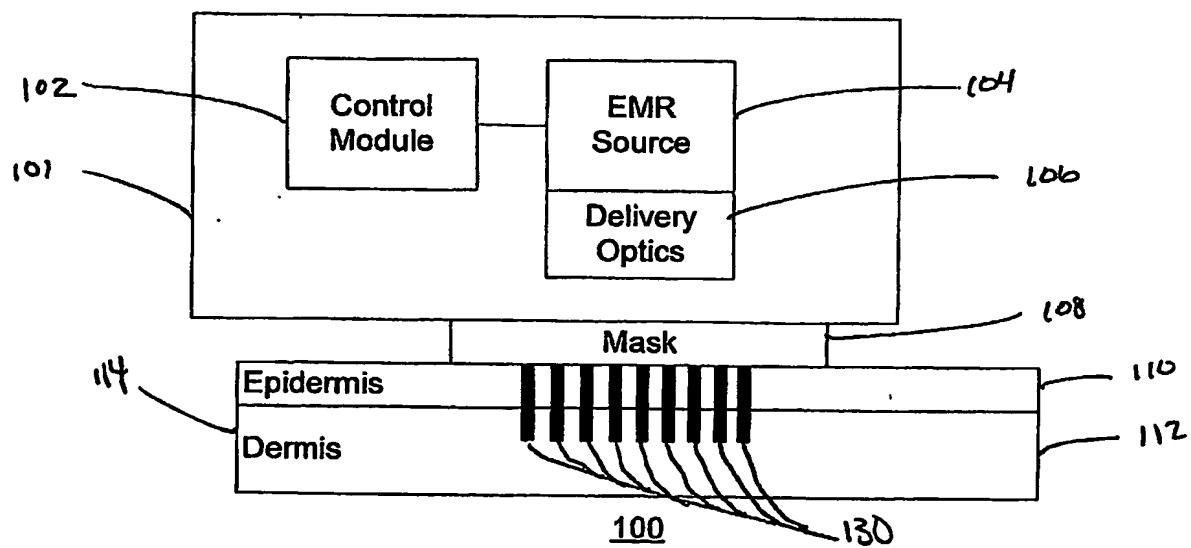
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100  
Fig. 1A



100  
Fig. 1B



100  
Fig. 1C

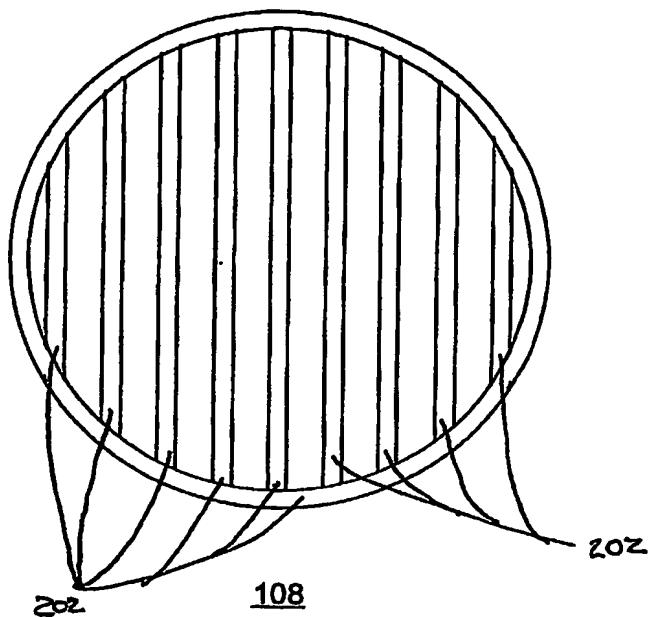
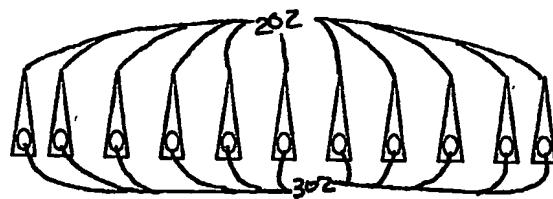
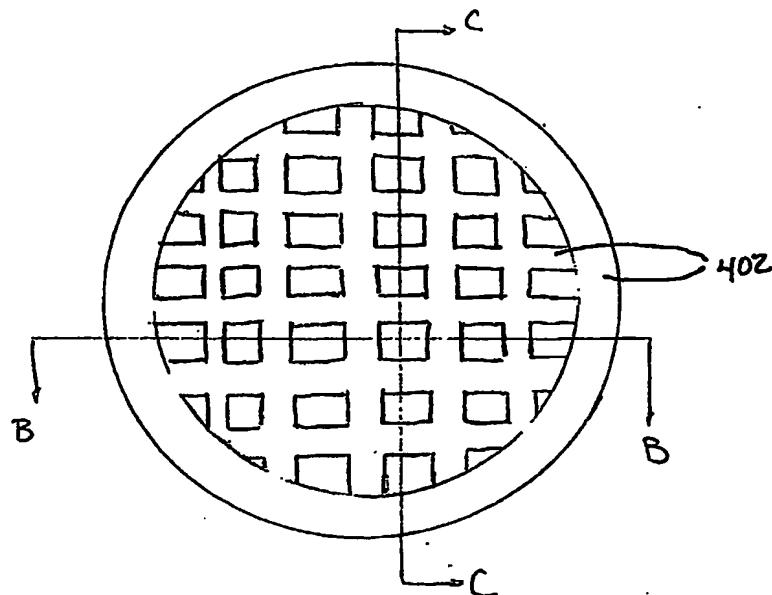


Fig. 2



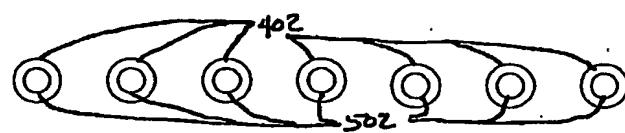
A-A  
Fig. 3



400

Fig. 4

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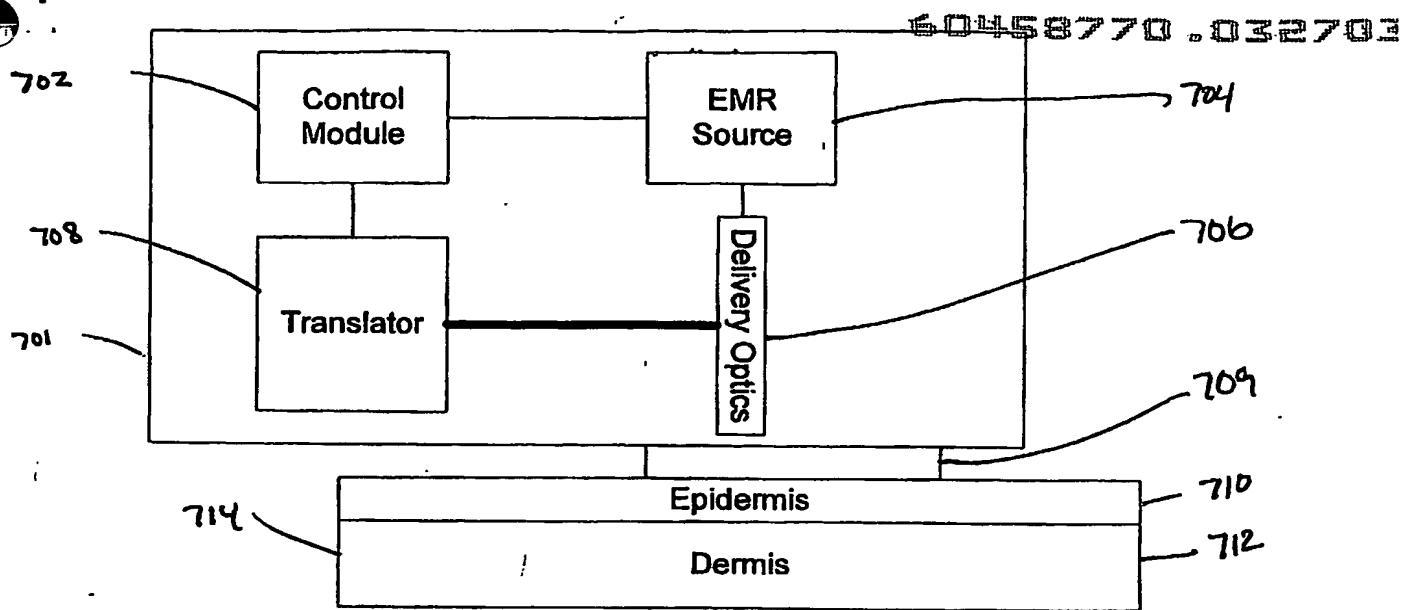
B-B

Fig. 5

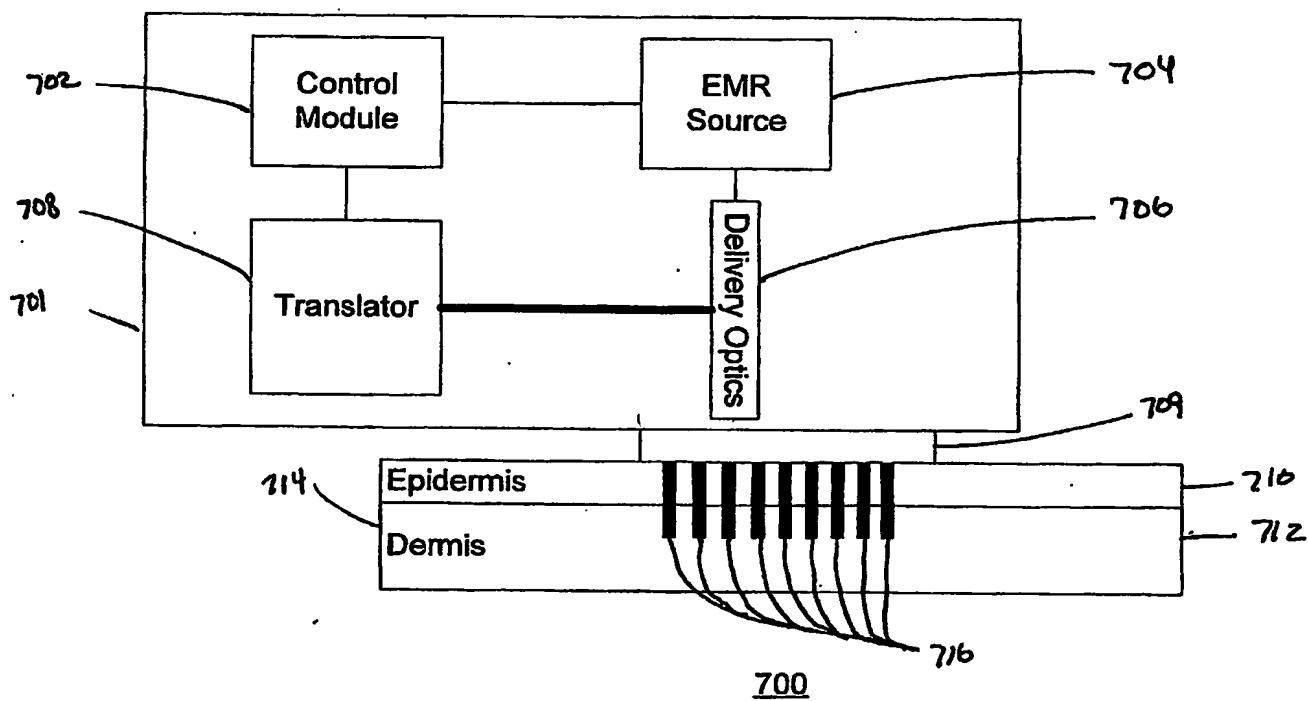


C-C

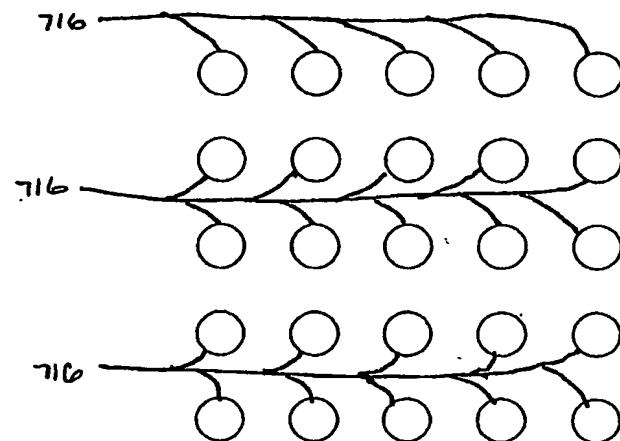
Fig. 6



700  
Fig. 7A

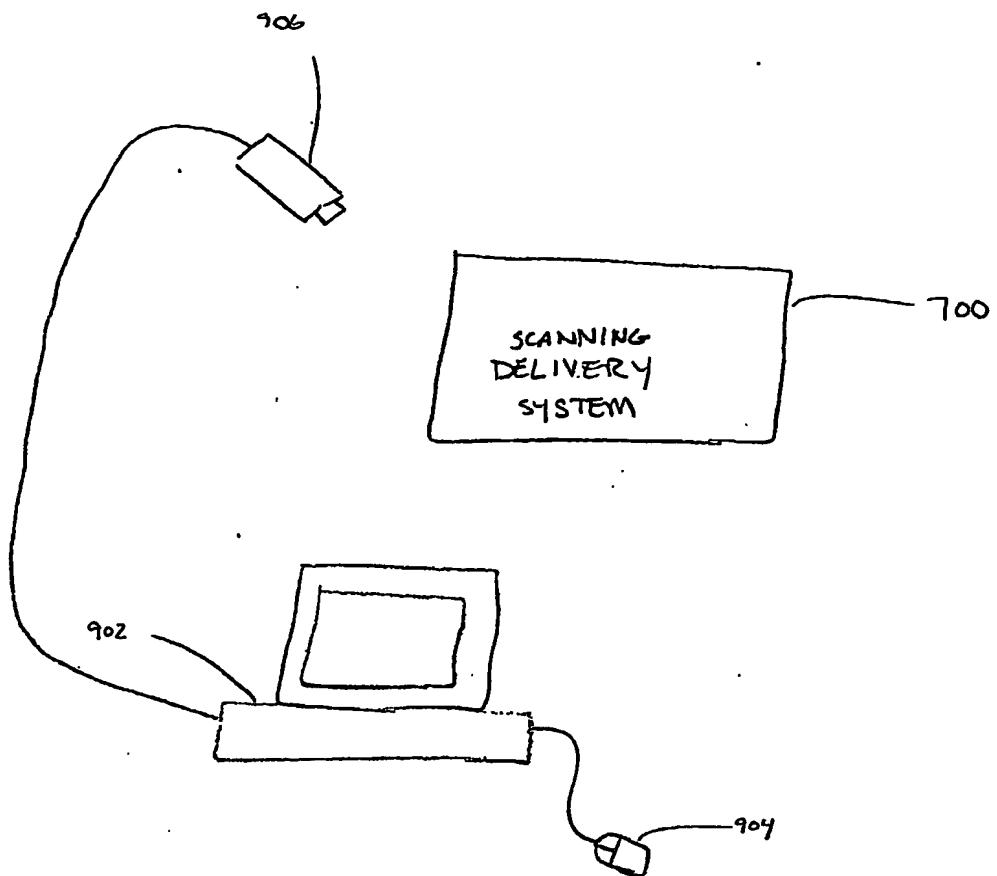


700  
Fig. 7B



800

Fig. 8



900

Fig. 9